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Huvudfaxen Kassan

TITLE

ANTIBACTERIAL MATERIAL

DESCRIPTION 5

Technical field

The present invention relates to an anti-bacterial material to be used for removing micro organisms such as bacteria, fungi or virus from surfaces, such as organic surfaces including skin and including wounds, construction surfaces including building surfaces, furniture surfaces and automotive surfaces, as well as for removing microbes in air by being present in a filter form.

The object of the present invention is to obtain an anti-microbial material, which can be used for primarily removing bacteria, fungl and/or virus from different surfaces and 15 substrates without wetting or contaminating the surface with different antibacterial agents.

A second object is to obtain a material, which has a microbicidal effect as well.

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Background of the invention

It is generally known to prepare materials of different structures and thereby providing them with a liquid, which may contain an anti-bacterial effect as it may contain an antibacteriostatic or bactericidal substance.

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Quite often the liquid is an alcohol, such as ethanol or isopropanol, which in itself is bactericidal, or at least bacteriostatic.

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The problem is the wiping around with such a material, which will contaminate surfaces around the primary target to be cleansed, and complete removal of bacteria will thus not be obtained.

The use of wet materials is not always appropriate as the liquid may damage the surface treated, a treated wound may be hurting from concentrated alcohol or even damages may occur due to sensitivity.

US 6,258,455 discloses an anti-microbial ultra-microfilament cloth having one yarn of fine fiber of 1.0 denier or less, and one yarn having an antimicrobial fiber, which yarns

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are engaged with each other. The antimicrobial fiber consists of a cellulose fiber having Havudföxen kassan been acetate treated to obtain antimicrobial properties. The problem solved is to prevent propagation of bacteria and fungi in cleansing cloths as common cloths just remove such microbes. From a general standpoint of view it is indicated that such cloths just spread the microbes around, and do not absorb them. By having an antimicrobial effect such microbes should be killed.

An ultra microfiber consisting of polyamide and polyester is said to be superior conventional fibers due to its finess and due to its triangular cross-section. The cross-section provides for a mechanical lifting off of the microbes. It is also said to have a positive charge for attracting the microbes, which charge is due to the polyamide. The fiber will, however, provide a possibility for continued propagation of the microbes. To attain antimicrobial activity triclosan is embedded in the acetate fiber, whereby triclosan has an ability of penetrating the cell walls of thin walled cells.

EP-A-1,091,676 discloses a skin care device with a cleansing tool comprising a padding configuration consisting of at least in part of microfiber tissue. Thus the cleansing effect is related to the microfiber as such.

20 DE-A-19,839,505 relates to a cleansing cloth being a knitted fabric with woven zones providing a cleansing surface with projecting microfiber loops or piles produced in an automatic production line.

EP-A-0 286 741 relates to a polyamide yarn provided with built-in antibacterial capacity whereby an adhesive on the fiber surface of an antimicrobial agent consisting of an organosilicon, quaternary ammonium salt compound and a surfactant is present thereon.

WO 01/96433 relates to the use of polymers containing urethane and/or urea groups for the modification of surfaces, whereby the polymers are used on particulate materials, line-shaped, planiform or three-dimensional entities.

JP 200003284 relates to surface modification of antibacterial agents for fibers to be able to incorporate the antimicrobial agent in the fiber without causing problem to the fiber when knitted, woven or otherwise used.

US-A-4,847,089 relates to a cleansing and disinfecting composition comprising an alkaline per-salt and a positively charged phase-transfer agent and absorbent materials,

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such as sponges, impregnated therewith. The positively charged phase-transfer agent is preferably a quaternary ammonium salt. The composition may be formulated as a cream, a bulk powder, a solution or tablets. It may also be incorporated in wipes, sponges and brushes. In one embodiment, one surface of the impregnated sponge is an abrasive material. The compositions of the invention according to USP 4,847,089 find utility in health care, as disinfectants in surgical applications, and as cleansing and decontaminating agents. Specific uses include skin cleansing and disinfecting.

In the cleaning of surfaces including skin, other soft surfaces, as well as hard surfaces it is often of utmost interest and importance to remove microbes in form of bacteria and fungi or viruses by mechanical means and/or by using an antimicrobial agent that kills or makes the microbes passive. The bacteria of interest to become removed are Escherichia coli, Staphylococcus aureus and different Streptococcus strains. E. coli may give raise to different gastro-intestinal illnesses, while S. aureus may cause troublesome infections, like certain Streptococci.

It is also known that most cells are electrically charged and this has been the basis of the present invention.

20 Summary of the present invention

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The present invention thus primarily relates to a antimicrobial material for absorbing and retaining microbes, including bacteria, fungl and/or virus, characterized in that it has a positive charge that creates an electrical field strength of at least 100 V/m, preferably the positive charge creates an electrical field strength of at least 150 V/m, more preferably the positive charge creates an electrical field strength of at least 200 V/m.

In a preferred embodiment the material consists of a fibrous, non-woven material.

30 In a further preferred embodiment the material consists of a fibrous, woven material.

In another preferred embodiment the material consists of a naturally occurring fiber.

In a further preferred embodiment the material consists of an inorganic fiber.

In another preferred embodiment the material consists of an organic, polymeric fiber.

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In a further preferred embodiment the surface consists of an organic polyment -0.1- 1.4 composite fiber,

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In another preferred embodiment the material is in the form of a layered structure.

In another further preferred embodiment the material is in the form of a layered structure having a second treating ingredient(-s) on a side opposite to the positively charged side.

In a preferred embodiment the inventive material is to be used in health care, including hospital care.

Detalled description of the present invention

By identifying the specific surface charge being antimicrobial the invention is independent of chemistry and chemical processes, but the invention is based on electrostatic interaction between a charged surface and the surface of microbial cell thereby providing an antimicrobial effect. This makes it possible to produce a number of chemically varied entities and manufacturing processes taking into account the specific application of the product prepared.

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By using electrostatic interaction together with cell surfaces a number of different intracellular as well as extracellular processes are generated, such as electrostatic attraction with intracellular charges and the binding of a microbe onto a charged surface, mediation of transmembrane ion channels, and mediation of metabotrophic receptors and intracellular signal pathways, as well as other routes of action.

The level of electrostatic charge to attract microbes was primarily determined in a series of test.

30 Test 1.

E. coli and S. aureus strains were used in the test to be described below. The bacteria, one type in each test, are suspended in a physiological saline solution (0.9 % NaCl). Glass substrates were covered with aluminium foil to be able to apply an electrical field over the bacterial suspension applied to the substrate. The substrate and aluminium foil are connected to an electric source having constant-regulated voltage, and is brought under a light microscope for analysis. Different voltages in the range of 0 to 40 V are applied. The movements of the bacteria suspended are studied. In order to differentiate between filed dependent movements and other types of movements the polarity is

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changed. Different voltages were applied over the substrate. The following pasultspyere 4 obtained.

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	Voltage	Observations
5	0-3 V	No influence on the movements of the bacteria. However, a
		movement perpendicular to the electrical field was observed,
		which probably was due to an increase of temperature.
	4 V	Evident threshold level of influencing the movements of
		bacteria parallel to the electrical field lines In spite of polarity
10		changes the bacteria strive towards the positive pole.
	5-40 V	As for 4 V but more powerful movements. Even perpendicular
		movements increase.

The field strength at the point observed depends on the geometry of the observed position on the substrate relative the electricity conducting Al-poles. The field strength estimated at the point under observation was about 200 V/m at 4 V. Field strengths in the range of 100 V/m have an evident threshold effect in this test. The relative dielectricity constant of the solution was not considered.

20 It was observed that there was no difference between the bacteria strains tested.

Test 2.

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Further tests were conducted using treated surfaces to provide for an electrical charge great enough to attract microbes. Chemically modified surfaces consisting of glass fiber filters and sterile plasters were used. The glass fiber substrate and the sterile plasters were modified by spreading 1 mi of 3-aminopropyl-triethoxy silane over the material surface for 5 min. The substrates were then placed in a heating cabinet to obtain a reaction between the EtO-Si-groups of the chemical and M-OH groups of the substrates. The chemically modified surfaces were exposed for 10 s to S. aureus and diphtheroidal rods in a nutrient solution (PBS). A cautious drainage took place for 20 s whereupon the substrates were rapidly pressed against a cultivation plate (agar). Cultivation took place in a heating cabinet for 24 hrs. The samples were read and photographed. Cultivation from the modified surfaces was 40 % lower than cultivation from non-modified surfaces.

35 Test 3.

Slx glass fiber surfaces were modified in the same way as described in Test 2. The different surfaces differ from each other with regard to surface charge and type of modification. The chemicals were used in relative concentrations 1:1, 1:10, and 1:50.

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The surface charge density for a dilution 1:1 can be expressed as if 2 ml 3-philipping triestor triethoxy silane are attached to a macroscopic surface of 1 dm². The active surface is, however, probably considerably larger due to the surface enlargement obtained by the fiber structure of the material used.

The chemistry differed with regard to the outer end of the molecule. In the first group cases the molecule was ended with $-N^+(CH_3)_3$, and in the second group cases (Test 3) with $-NH_2$ groups. A substantial difference is thus the charge at neutral pH.

The tests have been carried out using 3-(trimethylammonium)propyl triethoxy silane and 3-aminopropyl-triethoxy silane, but also other substances can be used for obtaining a positive charge, such as other compounds attachable to the substrate and containing a free amino group, or a substituted amino group.

Other compounds, which have turned out feasible, are silicon compounds being cationicly charged.

Further, compositions of the invention are based on

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[3-(trimethoxysllyI)]propyldimethyloctadecylammonium chloride), $C_{26}H_{58}CINO_3Si$, CAS number: 27668-52-6,

which can be applied onto fibers, fabrics, cloths, etc of polyester, cotton, glass, etc.

To improve, increase, hydrophilicity of said fibers, fabrics, cloths, etc treated with said octadecylammonium chloride the coating can be treated with

A) N-trimethoxysilylpropyl-N,N,N-trimethylammonium chloride

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C9H24CINO3SI

CAS number: 35141-36-7

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or

B) Polyethyleneimine, CAS number: 25987-06-8

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Experimental procedures for coating of polyester/polyamide cloth

10% of the octadecylammonium chloride compound above: The [3-(trimethoxysilyl)]propyldimethyloctadecylammonium chloride) is diluted 10 % by weight in water and is stirred for 15 min at room temperature.

10 % of N-trimethoxysilylpropyl-N,N,N-trimethylammonium chloride. The N-trimethoxysilylpropyl-N,N,N-trimethylammonium chloride is diluted to 10 % by weight in water and is stirred for 30 min at room temperature.

5% of polyethylene imine: polyethylene imine is dijuted to 5 % by weight in water and is stirred for 30 min at room temperature.

25 10% of an 1:1 solution of [3-(trimethoxysilyl)]propyldimethyloctadecylammonium chloride)/ N-trimethoxysilylpropyl-N,N,N-trimethylammonium chloride: [3-(trimethoxysilyl)]propyldimethyloctadecylammonium chloride) and N-trimethoxysilylpropyl-N,N,N-trimethylammonium chloride are mixed in equal proportions and are diluted to 10 % by weight by adding water and stirring for 15 min at room temperature.

Coating using [3-(trimethoxysilyi)]propyldimethyloctadecylammonium chloride) and polyethylene imine

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The cloth is dipped for 1 min in the 10 % solution of [3(trimethoxysilyi)]propyldimethyl octadecylammonium chloride) and is the Handistated Cassan
The cloth is then heated to 150oC for 2 minutes. Yhereafter the cloth is dipped into a
solution of 5 % by weight of polyethylene imine for 1 minute and is then squeezed and
dried.

Coating using an 1:1 solution of above.

The cloth is dipped into a 1:1 10% solution of [3-

10 (trimethoxysilyl)]propyldimethyloctadecylammonium chloride)/ N-trimethoxysilylpropyl-N,N,N-trimethylammonium chloride for 1 minute and is then squeezed and dried at 150oC for two minutes

The different surfaces were exposed to a bacterium suspension of £. coli for 10 s.

Drainage of surplus amount of liquid was carried out, whereupon the surfaces were rinsed with physiological saline solution for 20 s. The surfaces were allowed to dry before an electron microscopy examination took place. The result was somewhat difficult to read due to the drying as the cells dry out and their three-dimensional structure get lost. Certain detection is made in the form of traces after cell walls. Single cells have their structure left.

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A second test was made whereby no drying was carried out. Electron microscopy 1-14 examination was carried out immediately. Felts A, B. and C were all treated using Kusson (trimethylammonium) propyl triethoxy silane, and Wovens A, B, and C were treated with 3-aminopropyl triethoxy silaneThe results obtained are given below.

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	Substrate	Conc.	Observation I	Observation II
	Felt A	1:10	Only single bacteria cells	
			can be detected. "Prints"	
			of dried cell walls are,	
10			however, noticed.	
	Felt B	1:1	As above	
	Felt C	1:50		A large amount of bacteria
				was observed.
	Felt, neg1)			Only single cells can be detected
15	Woven A	1:10	Only single cells can	
			be detected	
	Woven B	1:1	Only single cells can	
			be detected.	
	Woven C	1:50	Only single cells can	
20			be detected.	
	Woven neg1)	Only single cells can	
			be detected.	

¹⁾ neg means untreated clean surface used as reference.

The results obtained provide for the following understanding. In Observation I residues of bacteria are seen on all felt surfaces. No significant difference is seen, however, between the different preparations. This can be interpreted as concentration differences at the preparation of the surface i) does not give raise to significant differences in the ability of attracting bacteria, ii) that the preparations in spite of different handling does not significantly differ with regard to surface structure/charge density, or iii) that the concentration differences are incorrectly chosen with too small differences, or in the wrong range, alternatively.

Observation II gave a significant difference between Felt C and Felt neg which result is interpreted as an already low concentration difference at the preparation of the surfaces provides for surface characteristics enough to differ from an untreated surface. This observation is central for the interpretation of the results.



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When It comes to the woven structures there are no significant differences noticed between samples A, B, C, and neg., respectively. The modifications of these structures thus have a low influence on the ability of attracting bacteria. This is explained by the uncharged surface structure. Structurally, these surfaces, however, show great similarity with the surfaces of the felt pieces.

The surfaces to be coated are any non-woven or woven structure of any natural structure, such as cellulose, cotton, wool, or other naturally occurring fibres or mixtures thereof, inorganic fibers, polymeric or mixed or otherwise composite fiber material in non-layered or layered, including multilayered form, as well as pure fiber web structures used for e.g., filtering purposes. Cellulose fibers are often used in non-woven tissue, even paper, like inorganic fibers, such as glass and stone fibers or fibrous calcium sulphate. Other naturally occurring fibers are often used as filaments in woven materials. Polymeric fibers and composite fibers, such as carbon fiber, polyamide fiber, polyolefin fiber, polyester fiber, which can be used in a non-woven structure as well as in a woven structure.

In case of layered products one side thereof may be provided with a positively charged side that creates an electrical field strength of at least 100 V/m, while the other side comprises e.g., a wound cleansing preparation in the form of a saline solution, a wound healing preparation in the form of an ointment or more or less viscous liquid, or gel, possibly containing nourishing substances and/or growth factors, a skin lotion, a sun protecting lotion, a skin moisturing lotion, a skin "tanning" composition.

Other applications are as filtering aids in breathing masks or venting systems in prenatal incubators, buildings or vehicles, such as rail wagons, and cars, including trucks, where the fibers used for filtering the air are prepared for absorbing bacteria or virus, thereby eliminating access to the respiratory tract, e.g., in face masks for avoiding contamination of SARS, severe acute respiratory syndrome, virus or other types of viruses, such as influenza viruses. It has also been proposed for cleaning if possibly infected water, such as under field conditions.

Storage, either long-term or short-term, of foodstuffs shows different problems and the proliferation of different bacteria jeopardizes durability as well as consumer health. Great demands are put on storage at the right temperature and careful hygienic conditions to prevent rise, proliferation and the spread of bacteria. Since long it is well known that moisture and oxygen, as well as elevated temperatures nourish bacteria,

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virus and fungi. One application of the present invention aims to function individual moist environments.

Further applications are thus the use of a non-woven cloth as liquid absorbing material in food packages, such as meat packages, shellfish packages, fish packages, fowl and poultry packages, vegetable packages, fruit packages, such as trough of polymer materials, or cellulose material, wrappings, either of polymer or cellulose material, storage tinplates, and in general all types of material which can be provided with a positive charge, and can be used in or at the treatment of foodstuffs in general, a.o..

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Thus the invention can be used by the food industry, package industry, raising industry, culturing industry, transport industry, shipping, air transport industry, agricultural sector, slaughter-houses, largescale households, catering, kitchens, restaurants, a.o..

The most common bacteria to eliminate are Bacillus cereus, calivirus and virus similar to the Norwalk virus, including Norwalk, Campylobacteria, Clostridium botulinum, C. perfringens, Listeria monocytogenes, Penicillium, Aspergillus and Fusarium strains which give rise to mould, Salmonella strains, and Shigelia, Staphylococcus, and Yersinia strains. Certain bacteria give rise to production of histamine in the foodstuff, and histamine may cause changes in taste as well as more severe sicknesses.

Other applications are as surface cleansing cloths for hard or semi hard surfaces such as furniture, building walls and floors, sanitary equipment, and other equipment to prevent bacterial, fungal, or viral contamination. Thus so called miracle fibre cloths having a fibre of a particular triangular cross-section can be modified by positively charging the same in accordance with the present invention.

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Other applications are stockings and socks to reduce bacterial and fungi growth in "hot" feet. Other applications are as incontinence diapers, napkins, handkerchiefs, wound compresses, plaster, surgical cloths, surgery clothing, and pre-surgery coverings. In applications concerning incontinence diapers the primary aim is to take care of odour-producing bacteria, which quite often occur in such situations. The material of the invention is also well suited for use under field conditions, as there is no volatile liquid present.

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Surgical cloths can be used not only at surgery but also at the treatment of eczema and burn injuries, where the skin is sensitive to inflammation of microbes, in particular bacteria. Thus such cloths can be used for covering and/or sheets.

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In all applications it is also a question of obtaining a microbicidal effect, i.e., Howilling of Kassa any microbe attracted to the antimicrobial material of the invention.

The transfer of contagions, based on bacteria or viruses, respectively, can take place by drip contamination (contact) or air born contamination. Today, sterile gloves are used to prevent transfer of contagions at contact. However, using this method the gloves are being contaminated and a contamination will take place to any surface as the contaminated gloves will be brought into contact with. The method is, with other words, only applicable when one operates within a small area and the number of touched surfaces can be held to a minimum and these can be held under control. An alternative that kills any contagion and thereby eliminates contaminations is to treat the skin of the hands using an active chemical agent. This is, however, not recommended at long term use.

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A glove can be prepared in accordance with the present invention and its fibers are then being surface modified so that any contagion (bacteria or virus) is absorbed and killed in the material. Even if there should be no total killing of the contagions the innovation provide the user an improved freedom to move around compared to other methods, and one need not to worry about contact contamination or to spread the same. The solution further provides the user the possibility of obtaining an efficient protection without need for handling a number of chemical agents.

Generally the skin functions as an excellent barrier against contagions, whereby however, openings therein, such as a wound, will function as an opening of this barrier. With other words wounds means that the body will be open to contagions at such places. An alternative for creating an temporary barrier may be the use of a tight plaster covering the wound. The problem using this solution is the absence of air which normally prolongs the healing time. Another solution is to have an antibacterial plaster which prevents bacterial growth in the wound, i.e., in the environment closest to the wound surface. The problem of this solution is that some bacteria, which have a positive effect on the healing also will become destroyed.

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To sum up, one may state that open wounds are vulnerable to contagions. Several factors are necessary to have a rapid healing process being as fast as possible, but, with regard hereto it should be noted that on one side wounds need to be protected from bacteria coming from the outside, and on the other side wounds need air to provide for an optimal healing process.

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A barrier having antibacterial properties and which is simultaneously permeable to air is a solution to the problem. The plaster consists of two layers; the layer closest to the wound being an untreated material, such as an ordinary wound compress; the layer facing outwards is of a similar material but being surface modified using the present invention. Further, the outer surface is made extremely hydrophobic to provide for a protection against dirt and moisture which otherwise may penetrate into a wound. This solution provides air to the healing process simultaneously as the product is a bacterial barrier to contagions appearing from the surroundings, without creating an antibacterial atmosphere closest to the wound.

Traditional breath masks and mouth masks are based on the principle that a filtering material filtrates off particles and bacteria to prevent contagions and pollutions to enter the body via the respiratory tract. This function is, in the best case, active as long as the filtering material is dry.

When exhaust air and saliva have moistened the filtering material the antibacterial properties of the filtering material will be gone partly or completely. At normal use this will take place already after 7 to 15 minutes.

By covalent surface modification according to the present invention the filtering material of a breath or mouth mask obtains unique antibacterial properties functioning both in dry as well as wet surroundings. The invention solves these known problems of a conventional breath mask successively becomes ineffective and that the antibacterial properties thereby are eliminated.

The antimicrobial material of the invention can thus be used as filter, as cleansing cloth in health care including hospital care and medical treatment, in cosmetics, in general sanitarian care and maintenance. One example of a cleansing material is a dry cloth to be used by doctors, and nurses, or other personal in contact with different patients or patient groups, where there is a request for antibacterial cleansing between any new meeting with a patient, and where hitherto used alcoholic solutions tend to dry out the skin causing irritation and even supersensitivity.

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CLAIMS

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- 1. Antimicrobial material for absorbing and retaining microbes, including bacteria, fungi and/or virus,
 - characterized in
- that it has a positive charge that creates electrical field strength of at least 100 V/m.
 - 2. Antimicrobial material according to claim 1, wherein the positive charge creates electrical field strength of at least 150 V/m.
 - 3. Antimicrobial material according to claim 2, wherein the positive charge creates electrical field strength of at least 200 V/m.
 - 4. Antimicrobial material according to one or more of the preceding claims, wherein the material consists of a fibrous, non-woven material.
 - Antimicrobial material according to one or more of claims 1-3, wherein the material consists of a fibrous, woven material.
- 20 6. Antimicrobial material according to claims 1-5, wherein the material consists of a naturally occurring fiber.
 - 7. Antimicrobial material according to claims 1-5, wherein the material consists of an inorganic fiber.
 - 8. Antimicrobial material according to claims 1-5, wherein the material consists of an organic, polymeric fiber.
 - Antimicrobial material according to claims 1-5, wherein the surface consists of an organic polymeric composite fiber.
 - 10. Antimicrobial material according to claims 1-9, wherein the material is in the form of a layered structure.
 - 11. Antimicrobial material according to claims 10, wherein the material is in the form of a layered structure having a second treating ingredient(-s) on a side opposite to the positively charged side.

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- 12. Antimicrobial material according to claims 1-11 to be used in health care, including hospital care.
- 13. Antimicrobial material according to claims 1-11 to be used in an air filter as a filtering mass.
 - 14. Antimicrobial material according to claims 1-11 to be used in a water cleaning arrangement.
- 15. Antimicrobial material according to claims 1-11 to be used in food storage.
 - 16. Antimicrobial material according to claim 15, to be used in food packages.
 - 17. Antimicrobial material according to claim 15, to be used in food storage spaces.
 - 18. Antimicrobial material according to claim 15, to be used in food package for meat package, shellfish package, fish package, fowl and poultry package, vegetable package, fruit package.
- 20 19. Antimicrobial material according to claim 15, wherein the package is a trough of polymer materials, or cellulose material, wrappings, either of polymer or cellulose material, storage tinplates a.o..
- 20. Antimicrobial material according to one or more of claims 1-11, wherein the material is manufactured as gloves to prevent the spreading of contact contagions.
 - 21. Antimicrobial material according to one or more of claims 1-11, wherein the material is manufactured as a breath mask.
 - 22. Antimicrobial material according to one or more of claims 1-11, wherein the material is provided as an outer layer of a wound plaster, having an inner surface of a non-treated compress.
 - 23. Antimicrobial material according to one or more of the preceding claims, wherein the antimicrobial agent is [3- (trimethoxysilyi)]propyldimethyloctadecylammonium chloride, or another suitable salt thereof.

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- 24. Antimicrobial material according to claim 23, wherein N-trimethoxysilylpropyl-N,N,N-trimethylammonium chloride or another salt thereof is added to provide hydrophilicity.
- 25. Antimicrobial material according to claim 23, wherein polyethylene imine is added to provide hydrophilicity.

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ABSTRACT

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The present invention relates to a antimicrobial material for absorbing and retaining microbes, including bacteria, fungi and/or virus, whereby it possesses a positive charge which crates electrical field strength of at least 100 V/m, as well as certain uses thereof.

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